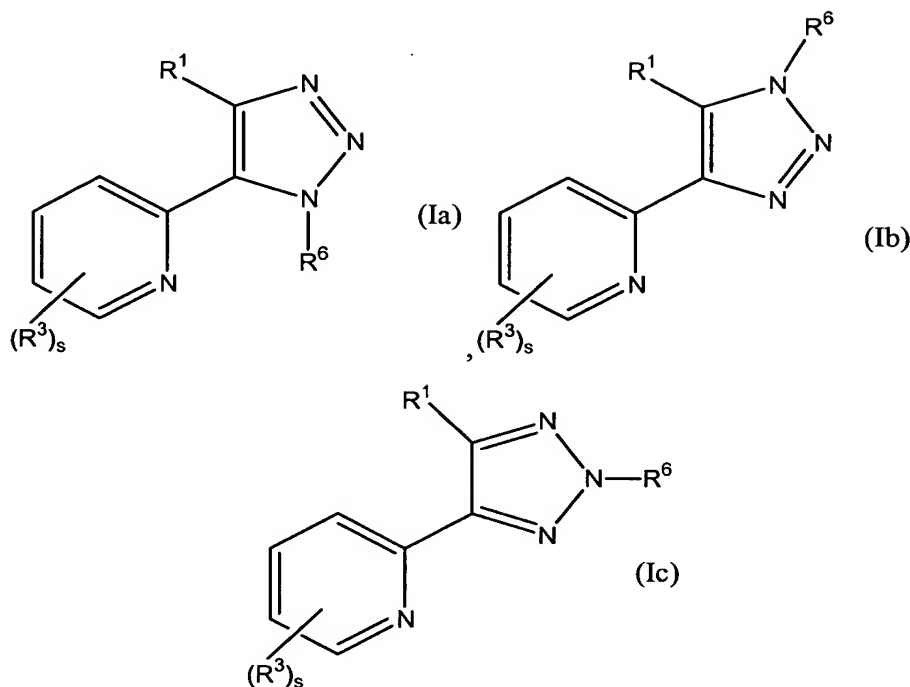


The claimed invention is:

1. A compound of formula (Ia), (Ib), or (Ic):



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or a pharmaceutically acceptable salt, prodrug, tautomer, hydrate or solvate thereof, wherein:

- $R^1$  is a saturated, unsaturated, or aromatic  $C_3$ - $C_{20}$  mono-, bi- or polycyclic ring optionally containing at least one heteroatom selected from the group consisting of N, O and S, wherein  $R^1$  can optionally be further independently substituted with
- 10 at least one moiety independently selected from the group consisting of: carbonyl, halo, halo( $C_1$ - $C_6$ )alkyl, perhalo( $C_1$ - $C_6$ )alkyl, perhalo( $C_1$ - $C_6$ )alkoxy, ( $C_1$ - $C_6$ )alkyl, ( $C_2$ - $C_6$ )alkenyl, ( $C_2$ - $C_6$ )alkynyl, hydroxy, oxo, mercapto, ( $C_1$ - $C_6$ )alkylthio, ( $C_1$ - $C_6$ )alkoxy, ( $C_5$ - $C_{10}$ )aryl or ( $C_5$ - $C_{10}$ )heteroaryl, ( $C_5$ - $C_{10}$ )aryloxy or
- 15 ( $C_5$ - $C_{10}$ )heteroaryloxy, ( $C_5$ - $C_{10}$ )ar( $C_1$ - $C_6$ )alkyl or ( $C_5$ - $C_{10}$ )heteroar( $C_1$ - $C_6$ )alkyl, ( $C_5$ - $C_{10}$ )ar( $C_1$ - $C_6$ )alkoxy or ( $C_5$ - $C_{10}$ )heteroar( $C_1$ - $C_6$ )alkoxy, HO-(C=O)-, ester, amido, ether, amino, amino( $C_1$ - $C_6$ )alkyl, ( $C_1$ - $C_6$ )alkylamino( $C_1$ - $C_6$ )alkyl, di( $C_1$ - $C_6$ )alkylamino( $C_1$ - $C_6$ )alkyl, ( $C_5$ - $C_{10}$ )heterocyclyl( $C_1$ - $C_6$ )alkyl, ( $C_1$ - $C_6$ )alkyl- and di( $C_1$ - $C_6$ )alkylamino, cyano, nitro, carbamoyl, ( $C_1$ - $C_6$ )alkylcarbonyl,

(C<sub>1</sub>-C<sub>6</sub>)alkoxycarbonyl, (C<sub>1</sub>-C<sub>6</sub>)alkylaminocarbonyl,  
 di(C<sub>1</sub>-C<sub>6</sub>)alkylaminocarbonyl, (C<sub>5</sub>-C<sub>10</sub>)arylcarbonyl, (C<sub>5</sub>-C<sub>10</sub>)aryloxycarbonyl,  
 (C<sub>1</sub>-C<sub>6</sub>)alkylsulfonyl, and (C<sub>5</sub>-C<sub>10</sub>)arylsulfonyl;

each R<sup>3</sup> is independently selected from the group consisting of: hydrogen,  
 5 halo, halo(C<sub>1</sub>-C<sub>6</sub>)alkyl, (C<sub>1</sub>-C<sub>6</sub>)alkyl, (C<sub>2</sub>-C<sub>6</sub>)alkenyl, (C<sub>2</sub>-C<sub>6</sub>)alkynyl,  
 perhalo(C<sub>1</sub>-C<sub>6</sub>)alkyl, phenyl, (C<sub>5</sub>-C<sub>10</sub>)heteroaryl, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic,  
 (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl, hydroxy, (C<sub>1</sub>-C<sub>6</sub>)alkoxy, perhalo(C<sub>1</sub>-C<sub>6</sub>)alkoxy, phenoxy,  
 (C<sub>5</sub>-C<sub>10</sub>)heteroaryl-O-, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic-O-, (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl-O-,  
 (C<sub>1</sub>-C<sub>6</sub>)alkyl-S-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-SO<sub>2</sub>-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-NH-SO<sub>2</sub>-, O<sub>2</sub>N-, NC-, amino,  
 10 Ph(CH<sub>2</sub>)<sub>1-6</sub>HN-, (C<sub>1</sub>-C<sub>6</sub>)alkyl HN-, (C<sub>1</sub>-C<sub>6</sub>)alkylamino, [(C<sub>1</sub>-C<sub>6</sub>)alkyl]<sub>2</sub>-amino,  
 (C<sub>1</sub>-C<sub>6</sub>)alkyl-SO<sub>2</sub>-NH-, amino(C=O)-, aminoO<sub>2</sub>S-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-(C=O)-NH-,  
 (C<sub>1</sub>-C<sub>6</sub>)alkyl-(C=O)-[((C<sub>1</sub>-C<sub>6</sub>)alkyl)-N]-, phenyl-(C=O)-NH-,  
 phenyl-(C=O)-[(C<sub>1</sub>-C<sub>6</sub>)alkyl)-N]-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-(C=O)-, phenyl-(C=O)-,  
 (C<sub>5</sub>-C<sub>10</sub>)heteroaryl-(C=O)-, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic-(C=O)-, (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl-(C=O)-,  
 15 HO-(C=O)-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-O-(C=O)-, H<sub>2</sub>N(C=O)-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-NH-(C=O)-,  
 [(C<sub>1</sub>-C<sub>6</sub>)alkyl]<sub>2</sub>-N-(C=O)-, phenyl-NH-(C=O)-, phenyl-[(C<sub>1</sub>-C<sub>6</sub>)alkyl)-N]-(C=O)-,  
 (C<sub>5</sub>-C<sub>10</sub>)heteroaryl-NH-(C=O)-, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic-NH-(C=O)-, (C<sub>3</sub>-  
 C<sub>10</sub>)cycloalkyl-NH-(C=O)- and (C<sub>1</sub>-C<sub>6</sub>)alkyl-(C=O)-O-;

20 where alkyl, alkenyl, alkynyl, phenyl, heteroaryl, heterocyclic, cycloalkyl,  
 alkoxy, phenoxy, amino of R<sup>3</sup> is optionally substituted by at least one substituent  
 independently selected from (C<sub>1</sub>-C<sub>6</sub>)alkyl, (C<sub>1</sub>-C<sub>6</sub>)alkoxy, halo(C<sub>1</sub>-C<sub>6</sub>)alkyl, halo,  
 H<sub>2</sub>N-, Ph(CH<sub>2</sub>)<sub>1-6</sub>HN-, and (C<sub>1</sub>-C<sub>6</sub>)alkylHN-;

25 s is an integer from one to five;

and

R<sup>6</sup> is selected from the group consisting of hydrogen, (C<sub>1</sub>-C<sub>6</sub>)alkyl,  
 30 (C<sub>2</sub>-C<sub>6</sub>)alkenyl, (C<sub>2</sub>-C<sub>6</sub>)alkynyl, phenyl, (C<sub>5</sub>-C<sub>10</sub>)heteroaryl, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic,  
 (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl, (C<sub>1</sub>-C<sub>6</sub>)alkyl-(SO<sub>2</sub>)-, phenyl-(SO<sub>2</sub>)-, H<sub>2</sub>N-(SO<sub>2</sub>)-,

- (C<sub>1</sub>-C<sub>6</sub>)alkyl-NH-(SO<sub>2</sub>)-, ((C<sub>1</sub>-C<sub>6</sub>)alkyl)<sub>2</sub>N-(SO<sub>2</sub>)-, phenyl-NH-(SO<sub>2</sub>)-, (phenyl)<sub>2</sub>N-(SO<sub>2</sub>)-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-(C=O)-, phenyl-(C=O)-, (C<sub>5</sub>-C<sub>10</sub>)heteroaryl-(C=O)-, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic-(C=O)-, (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl-(C=O)-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-O-(C=O)-, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic-O-(C=O)-, (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl-O-(C=O)-, H<sub>2</sub>N-(C=O)-,
- 5 (C<sub>1</sub>-C<sub>6</sub>)alkyl-NH-(C=O)-, phenyl-NH-(C=O)-, (C<sub>5</sub>-C<sub>10</sub>)heteroaryl-NH-(C=O)-, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic-NH-(C=O)-, (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl-NH-(C=O)-, ((C<sub>1</sub>-C<sub>6</sub>)alkyl)<sub>2</sub>N-(C=O)-, (phenyl)<sub>2</sub>N-(C=O)-, phenyl-(((C<sub>1</sub>-C<sub>6</sub>)alkyl)-N)-(C=O)-, (C<sub>5</sub>-C<sub>10</sub>)heteroaryl-(((C<sub>1</sub>-C<sub>6</sub>)alkyl)-N)-(C=O)-, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic-(((C<sub>1</sub>-C<sub>6</sub>)alkyl)-N)-(C=O)-, and
- 10 (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl-(((C<sub>1</sub>-C<sub>6</sub>)alkyl)-N)-(C=O)-;
- where alkyl, alkenyl, alkynyl, phenyl, benzyl, heteroaryl, heterocyclic, cycloalkyl, alkoxy, phenoxy, amino of R<sup>6</sup> is optionally substituted with at least one moiety independently selected from the group consisting of halo, (C<sub>1</sub>-C<sub>6</sub>)alkyl, (C<sub>2</sub>-C<sub>6</sub>)alkenyl, (C<sub>2</sub>-C<sub>6</sub>)alkynyl, perhalo(C<sub>1</sub>-C<sub>6</sub>)alkyl, (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl, phenyl,
- 15 benzyl, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic, (C<sub>5</sub>-C<sub>10</sub>)heteroaryl, (C<sub>1</sub>-C<sub>6</sub>)alkyl-SO<sub>2</sub>-, formyl, NC-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-(C=O)-, (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl-(C=O)-, phenyl-(C=O)-, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic-(C=O)-, (C<sub>5</sub>-C<sub>10</sub>)heteroaryl-(C=O)-, HO-(C=O)-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-O-(C=O)-, (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl-O-(C=O)-, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic-O-(C=O)-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-NH-(C=O)-,
- 20 (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl-NH-(C=O)-, phenyl-NH-(C=O)-, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic-NH-(C=O)-, (C<sub>5</sub>-C<sub>10</sub>)heteroaryl-NH-(C=O)-, ((C<sub>1</sub>-C<sub>6</sub>)alkyl)<sub>2</sub>N-(C=O)-, phenyl-(((C<sub>1</sub>-C<sub>6</sub>)alkyl)-N)-(C=O)-, hydroxy, (C<sub>1</sub>-C<sub>6</sub>)alkoxy, perhalo(C<sub>1</sub>-C<sub>6</sub>)alkoxy, (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl-O-, phenoxy, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic-O-, (C<sub>5</sub>-C<sub>10</sub>)heteroaryl-O-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-(C=O)-O-,
- 25 (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl-(C=O)-O-, phenyl-(C=O)-O-, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic-(C=O)-O-, (C<sub>5</sub>-C<sub>10</sub>)heteroaryl-(C=O)-O-, O<sub>2</sub>N-, amino, (C<sub>1</sub>-C<sub>6</sub>)alkylamino, ((C<sub>1</sub>-C<sub>6</sub>)alkyl)<sub>2</sub>-amino, formamidyl, (C<sub>1</sub>-C<sub>6</sub>)alkyl-(C=O)-NH-, (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl-(C=O)-NH-, phenyl-(C=O)-NH-, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic-(C=O)-NH-, (C<sub>5</sub>-C<sub>10</sub>)heteroaryl-(C=O)-NH-,
- 30 (C<sub>1</sub>-C<sub>6</sub>)alkyl-(C=O)-[(((C<sub>1</sub>-C<sub>6</sub>)alkyl)-N]-, phenyl-(C=O)-[(((C<sub>1</sub>-C<sub>6</sub>)alkyl)-N]-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-SO<sub>2</sub>NH-, (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl-SO<sub>2</sub>NH-, phenyl-SO<sub>2</sub>NH-,

(C<sub>5</sub>-C<sub>10</sub>)heterocyclic-SO<sub>2</sub>NH- and (C<sub>5</sub>-C<sub>10</sub>)heteroaryl-SO<sub>2</sub>NH-;

wherein the phenyl or heteroaryl moiety of a R<sup>6</sup> substituent is optionally further substituted with at least one radical independently selected from the group consisting of halo, (C<sub>1</sub>-C<sub>6</sub>)alkyl, (C<sub>1</sub>-C<sub>6</sub>)alkoxy, perfluoro(C<sub>1</sub>-C<sub>6</sub>)alkyl and

5 perfluoro(C<sub>1</sub>-C<sub>6</sub>)alkoxy,

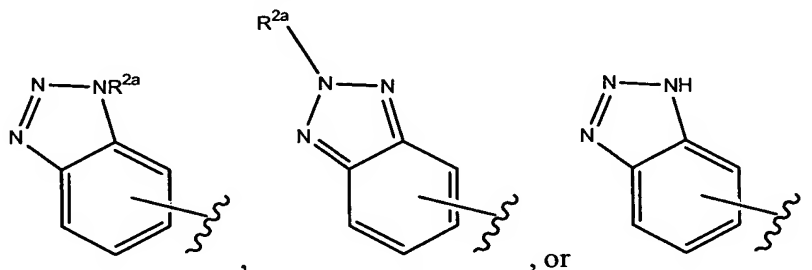
with the proviso that R<sup>1</sup> is not a naphthyl or phenyl; and

with the proviso that when R<sup>1</sup> is a phenyl fused with an aromatic or non-aromatic cyclic ring of 5-7 members containing up to three N atoms, said N is other than -NH or -NC<sub>1-6</sub>alkyl or if said N is -NH or -NC<sub>1-6</sub>alkyl, then R<sup>1</sup> must be further

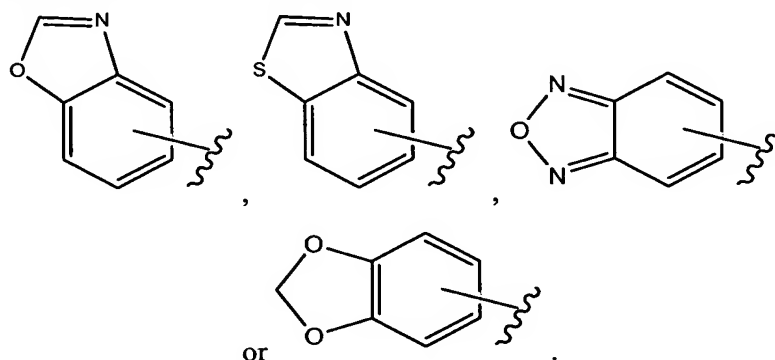
10 substituted; and

with the proviso that when R<sup>1</sup> is a phenyl fused with an aromatic or non-aromatic cyclic ring of 5-7 members containing 1-3 heteroatoms independently selected from O and S, then R<sup>1</sup> must be further substituted.

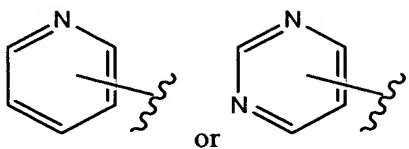
15 2. A compound of claim 1, wherein R<sup>1</sup> is



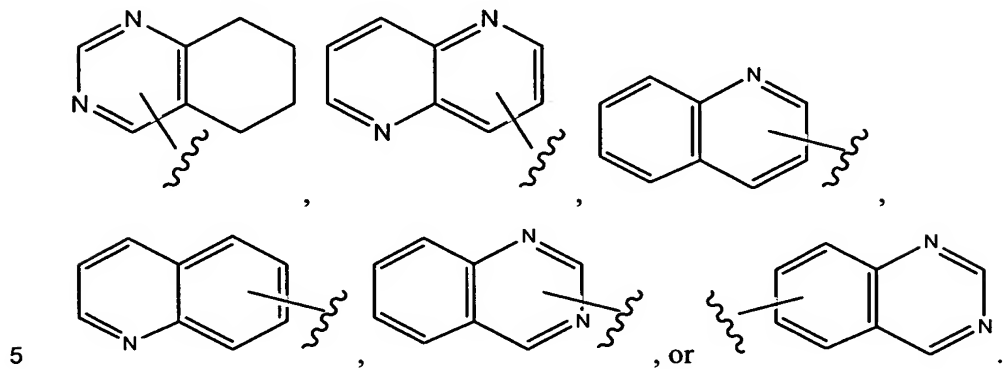
3. A compound of claim 1, wherein R<sup>1</sup> is



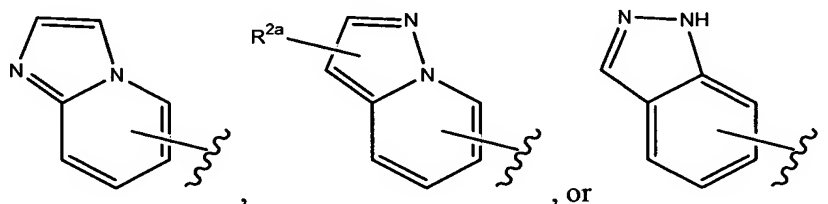
4. A compound of claim 1, wherein  $R^1$  is



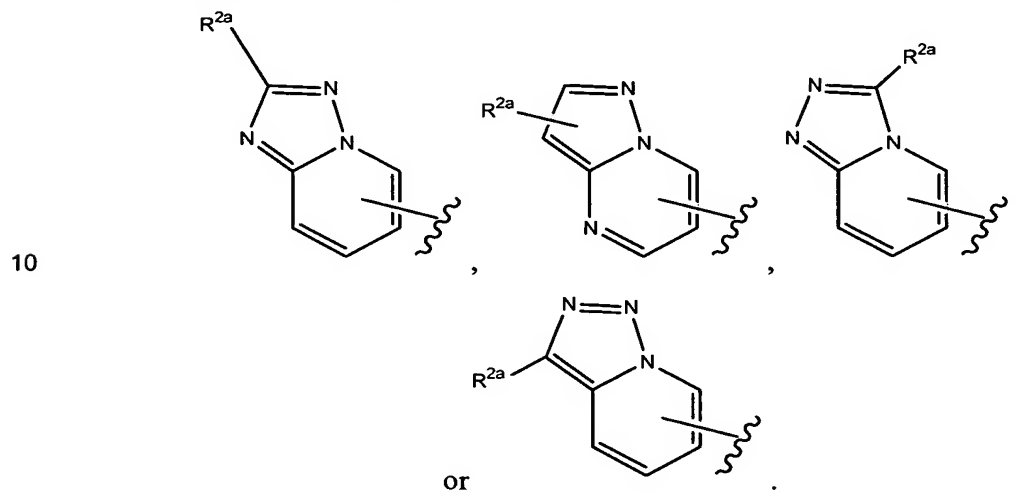
5. A compound of claim 1, wherein  $R^1$  is



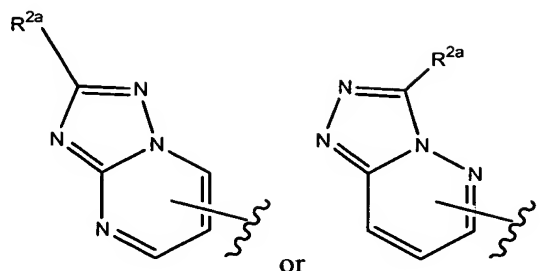
6. A compound of claim 1, wherein  $R^1$  is



7. A compound of claim 1, wherein  $R^1$  is



8. A compound of claim 1, wherein  $R^1$  is



9. A compound of claim 1, wherein  $s$  is one to two;  $R^3$  is hydrogen or  $(C_1-C_6)$ alkyl; and  $R^6$  is H,  $(C_1-C_6)$ alkyl, or  $(C_3-C_{10})$ cycloalkyl.

10. A pharmaceutical composition comprising a compound of claim 1 and a pharmaceutically acceptable carrier.

10

11. A method of preventing or treating a TGF-related disease state in an animal or human comprising the step of administering a therapeutically effective amount of a compound of claim 1 to the animal or human suffering from the TGF-related disease state.

15

12. A method of claim 11, wherein said TGF-related disease state is selected from the group consisting of cancer, glomerulonephritis, diabetic nephropathy, hepatic fibrosis, pulmonary fibrosis, intimal hyperplasia and restenosis, scleroderma, and dermal scarring.

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